

EXHIBIT B
CLEAN VERSION OF THE PENDING CLAIMS

1. An isolated nucleic acid molecule comprising a nucleotide sequence selected from the group consisting of:
 - (a) the nucleotide sequence as set forth in SEQ ID NO:1 or SEQ ID NO:3;
 - (b) a nucleotide sequence encoding the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
 - (c) a nucleotide sequence which hybridizes under moderately or highly stringent conditions to the complement of (a) or (b), wherein the encoded polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4; and
 - (d) a nucleotide sequence complementary to any of (a)-(c).

2. An isolated nucleic acid molecule comprising a nucleotide sequence selected from the group consisting of:
 - (a) a nucleotide sequence encoding a polypeptide that is at least about 70, 75, 80, 85, 90, 95, 96, 97, 98, or 99 percent identical to the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
 - (b) a nucleotide sequence encoding an allelic variant or splice variant of the nucleotide sequence as set forth in SEQ ID NO:1 or SEQ ID NO:3, wherein the encoded polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
 - (c) a nucleotide sequence of SEQ ID NO:1 or SEQ ID NO:3; (a) or (b) encoding a polypeptide fragment of at least about 25 amino acid residues, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
 - (d) a nucleotide sequence of SEQ ID NO:1 or SEQ ID NO:3, or (a)-(c) comprising a fragment of at least about 16 nucleotides;
 - (e) a nucleotide sequence which hybridizes under moderately or highly stringent conditions to the complement of any of (a)-(d), wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4; and
 - (f) a nucleotide sequence complementary to any of (a)-(c).

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3. An isolated nucleic acid molecule comprising a nucleotide sequence selected from the group consisting of:

- (a) a nucleotide sequence encoding a polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4 with at least one conservative amino acid substitution, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
- (b) a nucleotide sequence encoding a polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4 with at least one amino acid insertion, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
- (c) a nucleotide sequence encoding a polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4 with at least one amino acid deletion, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
- (d) a nucleotide sequence encoding a polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4 which has a C- and/or - terminal truncation, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
- (e) a nucleotide sequence encoding a polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4 with at least one modification selected from the group consisting of amino acid substitutions, amino acid insertions, amino acid deletions, C-terminal truncation, and N-terminal truncation, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;
- (f) a nucleotide sequence of (a)-(e) comprising a fragment of at least about 16 nucleotides;
- (g) a nucleotide sequence which hybridizes under moderately or highly stringent conditions to the complement of any of (a)-(f), wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4; and
- (h) a nucleotide sequence complementary to any of (a)-(e).

4. (Amended) A vector comprising the nucleic acid molecule of Claim 1.

5. A host cell comprising the vector of Claim 4.

8. A process of producing a Cloaked-2 polypeptide comprising culturing the host cell

of Claim 5 under suitable conditions to express the polypeptide, and optionally isolating the polypeptide from the culture.

9. A polypeptide produced by the process of Claim 8.

10. The process of Claim 8, wherein the nucleic acid molecule comprises promoter DNA other than the promoter DNA for the native Cloaked-2 polypeptide operatively linked to the DNA encoding the Cloaked-2 polypeptide.

11. The isolated nucleic acid molecule according to Claim 2 wherein the percent identity is determined using a computer program selected from the group consisting of GAP, BLASTP, BLASTN, FASTA, BLASTA, BLASTX, BestFit, and the Smith-Waterman algorithm.

12. (Amended) A process for determining whether a compound inhibits Cloaked-2 polypeptide activity or production comprising exposing a cell according to Claim 5 to the compound, and measuring Cloaked-2 polypeptide activity or production in said host cell.

13. An isolated polypeptide comprising the amino acid sequence set forth in SEQ ID NO:2 or SEQ ID NO:4.

14. An isolated polypeptide comprising the amino acid sequence selected from the group consisting of:

(a) the mature amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4, optionally further comprising an amino-terminal methionine;

(b) an amino acid sequence for an ortholog of SEQ ID NO:2 or SEQ ID NO:4, wherein the encoded polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;

(c) an amino acid sequence that is at least about 70, 80, 85, 90, 95, 96, 97, 98, or 99 percent identical to the amino acid sequence of SEQ ID NO:2 or SEQ ID NO:4, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;

(d) a fragment of the amino acid sequence set forth in SEQ ID NO:2 or SEQ ID NO:4 comprising at least about 25 amino acid residues, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;

(e) an amino acid sequence for an allelic variant or splice variant of either the amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4, or at least one of (a)-(c) wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4.

15. An isolated polypeptide comprising the amino acid sequence selected from the group consisting of:

(a) the amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 with at least one conservative amino acid substitution, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;

(b) the amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 with at least one amino acid insertion, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;

(c) the amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 with at least one amino acid deletion, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4;

(d) the amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 which has a C- and/or N-terminal truncation, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4; and

(e) the amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4, with at least one modification selected from the group consisting of amino acid substitutions, amino acid insertions, amino acid deletions, C-terminal truncation, and N-terminal truncation, wherein the polypeptide has an activity of the polypeptide as set forth in SEQ ID NO:2 or SEQ ID NO:4.

16. A polypeptide according to claim 14 or 15 wherein the amino acid at position 9 of SEQ ID NO: 2 is aspartic acid or glutamic acid.

17. A polypeptide according to claim 14 or 15 wherein the amino acid at position 39 of SEQ ID NO: 2 is glycine, proline, or alanine.

18. A polypeptide according to claim 14 or 15 wherein the amino acid at position 58 of SEQ ID NO: 2 is arginine, lysine, glutamine, or asparagine.

19. A polypeptide according to claim 14 or 15 wherein the amino acid at position 81 of SEQ ID NO: 2 is valine, isoleucine, methionine, leucine, phenylalanine, alanine, or norleucine.

20. A polypeptide according to claim 14 or 15 wherein the amino acid at position 102 of SEQ ID NO: 2 is tryptophan, tyrosine, or phenylalanine.

21. A polypeptide according to claim 14 or 15 wherein the amino acid at position 154 of SEQ ID NO: 2 is serine, threonine, or alanine.

a3 22. (Amended) An isolated polypeptide encoded by the nucleic acid molecule of Claim 1.

23. The isolated polypeptide according to Claim 14 wherein the percent identity is determined using a computer program selected from the group consisting of GAP, BLASTP, BLASTN, FASTA, BLASTA, BLASTX, BestFit, and the Smith-Waterman algorithm.

24. An antibody or fragment thereof produced by immunizing an animal with a peptide comprising an amino acid sequence of SEQ ID NO:2 or SEQ ID NO:4.

a4 25. (Amended) An antibody or fragment thereof that specifically binds the polypeptide of any one of Claims 13, 14, or 15.

26. The antibody of Claim 25 that is a monoclonal antibody.

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27. A hybridoma that produces a monoclonal antibody that binds to a peptide comprising an amino acid sequence of SEQ ID NO:2 or SEQ ID NO:4.

28. (Amended) A method of detecting or quantitating the amount of Cloaked-2 polypeptide using the anti-Cloaked-2 antibody or fragment of any one of Claims 24, 25, or 26.

29. (Amended) A selective binding agent or fragment thereof that specifically binds at least one polypeptide wherein said polypeptide comprises the amino acid sequence selected from the group consisting of:

- (a) the amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4; and
- (b) a fragment of the amino acid sequence set forth in at least one of SEQ ID NO:2 or SEQ ID NO:4; and
- (c) a naturally occurring variant of (a) or (b).

30. The selective binding agent of Claim 29 that is an antibody or a fragment thereof.

31. The selective binding agent of Claim 29 that is a humanized antibody.

32. The selective binding agent of Claim 29 that is a human antibody or a fragment thereof.

33. The selective binding agent of Claim 29 that is a polyclonal antibody or a fragment thereof.

34. The selective binding agent of Claim 29 that is a monoclonal antibody or a fragment thereof.

35. The selective binding agent of Claim 29 that is a chimeric antibody or a fragment thereof.

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36. The selective binding agent of Claim 29 that is a CDR-grafted antibody or a fragment thereof.

38. The selective binding agent of Claim 29 which is a variable region fragment.

39. The variable region fragment of Claim 38 which is a Fab or a Fab' fragment.

40. A selective binding agent or fragment thereof comprising at least one complementarity-determining region with specificity for a polypeptide having the amino acid sequence of SEQ ID NO:2 or SEQ ID NO:4.

41. The selective binding agent of Claim 29 which is bound to a detectable label.

42. The selective binding agent of Claim 29 which antagonizes Cloaked-2 polypeptide biological activity.

43. A method for treating, preventing, or ameliorating a disease, condition, or disorder comprising administering to a patient an effective amount of a selective binding agent according to Claim 29.

44. A selective binding agent produced by immunizing an animal with a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:2 or SEQ ID NO:4.

Ab 45. (Amended) A hybridoma that produces a selective binding agent capable of binding a polypeptide according to any one of Claims 13, 14, or 15.

46. (Amended) A composition comprising the polypeptide of Claim 13 and a pharmaceutically acceptable formulation agent.

47. The composition of Claim 46 wherein the pharmaceutically acceptable

formulation agent is a carrier, adjuvant, solubilizer, stabilizer, or anti-oxidant.

48. The composition of Claim 46 wherein the polypeptide comprises the mature amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4.

a7 49. (Amended) A polypeptide comprising a derivative of the polypeptide of Claim 13.

50. The polypeptide of Claim 49 which is covalently modified with a water-soluble polymer.

51. The polypeptide of Claim 50 wherein the water-soluble polymer is selected from the group consisting of polyethylene glycol, monomethoxy-polyethylene glycol, dextran, cellulose, poly-(N-vinyl pyrrolidone) polyethylene glycol, propylene glycol homopolymers, polypropylene oxide/ethylene oxide co-polymers, polyoxyethylated polyols, and polyvinyl alcohol.

a8 52. (Amended) A composition comprising a nucleic acid molecule of Claim 1 and a pharmaceutically acceptable formulation agent.

53. A composition of Claim 52 wherein said nucleic acid molecule is contained in a viral vector.

a9 54. (Amended) A viral vector comprising a nucleic acid molecule of Claim 1.

55. (Amended) A fusion polypeptide comprising the polypeptide of Claim 13 fused to a heterologous amino acid sequence.

56. The fusion polypeptide of Claim 55 wherein the heterologous amino acid sequence is an IgG constant domain or a fragment thereof.

a10 57. (Amended) A method for treating, preventing or ameliorating a medical condition

comprising administering to a patient the polypeptide of Claim 13 or the polypeptide encoded by the nucleic acid of Claim 1.

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58. (Amended) A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:

(a) determining the presence or amount of expression of the polypeptide of Claim 13 or the polypeptide encoded by the nucleic acid molecule of Claim 1 in a sample; and

(b) diagnosing a pathological condition or a susceptibility to a pathological condition based on the presence or amount of expression of the polypeptide.

59. (Amended) A device, comprising:

(a) a membrane suitable for implantation; and

(b) cells encapsulated within said membrane, wherein said cells secrete a protein of Claim 13, and wherein said membrane is permeable to said protein and impermeable to materials detrimental to said cells.

60. (Amended) A method of identifying a compound which binds to a polypeptide comprising:

(a) contacting the polypeptide of Claim 13 with a compound; and

(b) determining the extent of binding of the polypeptide to the compound.

61. (Amended) A method of modulating levels of a polypeptide in an animal comprising administering to the animal the nucleic acid molecule of Claim 1.

62. (Amended) A transgenic non-human mammal comprising the nucleic acid molecule of Claim 1.

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